

We Claim:

1. A method for making particles from a liquid feed stock containing a pharmaceutical agent to produce particles suitable for pulmonary administration having a narrow particle size distribution comprising:
  - providing a feed stock comprising a pharmaceutically active agent and a solvent;
  - forcing said feed stock into a manifold defined between a vibratable element and a plate and forcing the feed stock through the plate, said plate comprising holes of at least one predetermined diameter, in order to produce droplets comprising a droplet size distribution wherein at least 80% of the droplets have a diameter within  $\pm 25\%$  of the median droplet diameter;
  - removing solvent from said droplets to produce particles suitable for pulmonary administration.
2. A method according to claim 1 further comprising vibrating said vibratable element in order to force said feed stock through the plate and produce droplets.
3. A method according to claim 2 wherein a piezoelectric element is coupled to said vibratable element.
4. A method according to claim 1 wherein said holes comprise a predetermined diameter of less than 30 microns.
5. A method according to claim 4 wherein said holes comprise a predetermined diameter of less than 10 microns.
6. A method according to claim 1 wherein said plate comprises holes having a first diameter of less than 30 microns and a second series of holes having a second diameter of  $\pm 50\%$  of said first diameter.
7. A method according to claim 6 wherein said second diameter is within  $\pm 20\%$  of said first diameter.

8. A method according to claim 7 wherein said first diameter is less than 10 microns.
9. A method according to claim 1 wherein said atomizer is provided with said feed stock at a feed rate of 5 ml/mn - 3500 ml/mn.
10. A method according to claim 1 wherein said particles are porous.
11. A method according to claim 1 wherein said particles comprise a MMD of less than 10 microns and a MMAD of 1 - 5 microns.
12. A method according to claim 1 wherein said particles comprise a particle size distribution wherein at least 90% of the particles have a diameter within a range of less than 4 microns.
13. A method according to claim 1 wherein at least 90% of the droplets have a diameter within  $\pm 25\%$  of the median droplet diameter.
14. A method according to claim 1 wherein at least 95% of the mass of the droplets have a diameter within  $\pm 25\%$  of the median droplet diameter.
15. A method according to any one of claims 1, 13, or 14 wherein the droplets have a diameter is within  $\pm 15\%$  of the median droplet diameter.
16. A method according to claim 15 wherein the droplets have a diameter within  $\pm 8\%$  of the median droplet diameter.
17. A method according to claim 1 wherein said solvent is removed by heating said droplets in a gas stream to produce dried particles.
18. A method according to claim 17 wherein said dried particles are collected.
19. A method for spray drying a feed stock containing a pharmaceutical agent to produce particles suitable for pulmonary administration having a narrow particle size distribution comprising:

providing a feed stock comprising a pharmaceutically active agent at a flow rate of at least 5 ml/min;

forcing said feed stock into a manifold defined between a vibratable element and a plate and forcing the feed stock through the plate, said plate comprising holes of at least one

5 predetermined diameter, in order to produce droplets;

drying said droplets in a gas stream to produce dried particles comprising a particle size distribution wherein at least 70% of the mass of the particles have a diameter within a 4 micron range; and

collecting said dried particles.

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20. A method according to claim 19 wherein the dried particles comprise a particle size distribution wherein at least 80% of the mass of the particles have a diameter within a 4 micron range.

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21. A method according to claim 19 wherein the dried particles comprise a particle size distribution wherein at least 90% of the mass of the particles have a diameter within a 4 micron range.

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22. A method according to any one of claims 19-21 wherein the dried particles have a diameter within a 3 micron range.

23. A method according to any one of claims 19-21 wherein the dried particles have a diameter within a 1.5 micron range.

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24. A method according to claim 19 further comprising vibrating said vibratable element in order to force said feed stock through the plate and produce droplets.

25. A method according to claim 24 wherein said plate is vibrated by coupling a piezoelectric element to said plate.

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26. A method according to claim 19 wherein said holes comprise a predetermined diameter of less than 30 microns.

27. A method according to claim 19 wherein said plate comprises holes having a first diameter of less than 30 microns and a second series of holes having a second diameter of  $\pm 50\%$  of said first diameter.

5 28. A method according to claim 27 wherein said second diameter is within  $\pm 20\%$  of said first diameter.

29. A method according to claim 28 wherein said first diameter is less than 10 microns.

10 30. A method according to claim 19 wherein said particles are porous.

31. A method according to claim 19 wherein said particles comprise a MMD less than 10 microns and a MMAD 1 – 5 microns.

11 12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 75 76 77 78 79 80 81 82 83 84 85 86 87 88 89 90 91 92 93 94 95 96 97 98 99 100